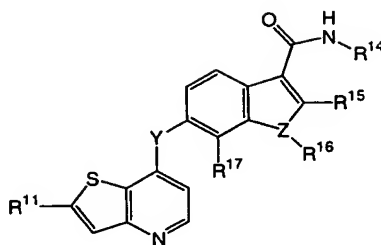


CLAIMS

We claim:

1. A compound represented by the formula I:



wherein:

Y is -NH-, -O-, -S-, or -CH₂-;

Z is -O-, -S-, or -N-;

10 R¹⁴ is a C₁-C₆ alkyl, C₁-C₆ alkylamino, C₁-C₆ alkylhydroxy, C₃-C₁₀ cycloalkylamino, or methylureido group;

R¹⁵ and R¹⁷ are independently H, halo, or a C₁-C₆ alkyl group unsubstituted or substituted by one or more R⁵ groups;

R¹⁶ is H or a C₁-C₆ alkyl group when Z is N, and R¹⁶ is absent when Z is -O- or -S-;

15 R¹¹ is H, C₁-C₆ alkyl, C₃-C₁₀ cycloalkyl, -C(O)NR¹²R¹³, -C(O)(C₆-C₁₀ aryl), -(CH₂)_i(C₆-C₁₀ aryl), -(CH₂)_i(5 to 10 membered heterocyclic), -(CH₂)_iNR¹²R¹³, -SO₂NR¹²R¹³ or -CO₂R¹², wherein said C₁-C₆ alkyl, -C(O)(C₆-C₁₀ aryl), -(CH₂)_i(C₆-C₁₀ aryl), and -(CH₂)_i(5 to 10 membered heterocyclic) moieties of the said R¹¹ groups are unsubstituted or substituted by one or more R⁵ groups;

20 each R⁵ is independently selected from halo, cyano, nitro, trifluoromethoxy, trifluoromethyl, azido, -C(O)R⁸, -C(O)OR⁸, -OC(O)R⁸, -OC(O)OR⁸, -NR⁶C(O)R⁷, -C(O)NR⁶R⁷, -NR⁶R⁷, -OR⁹, -SO₂NR⁶R⁷, C₁-C₆ alkyl, C₃-C₁₀ cycloalkyl, C₁-C₆ alkylamino, -(CH₂)_jO(CH₂)_qNR⁶R⁷, -(CH₂)_iO(CH₂)_qOR⁹, -(CH₂)_iOR⁹, -S(O)_i(C₁-C₆ alkyl), -(CH₂)_i(C₆-C₁₀ aryl), -(CH₂)_i(5 to 10 membered heterocyclic), -C(O)(CH₂)_i(C₆-C₁₀ aryl), -(CH₂)_iO(CH₂)_j(C₆-C₁₀ aryl), -(CH₂)_iO(CH₂)_q(5 to 10 membered heterocyclic), -C(O)(CH₂)_i(5 to 10 membered heterocyclic), -(CH₂)_jNR⁷(CH₂)_qNR⁶R⁷, -(CH₂)_jNR⁷CH₂C(O)NR⁶R⁷, -(CH₂)_jNR⁷(CH₂)_qNR⁹C(O)R⁸, (CH₂)_jNR⁷(CH₂)_iO(CH₂)_qOR⁹, -(CH₂)_jNR⁷(CH₂)_qS(O)_i(C₁-C₆ alkyl), -(CH₂)_jNR⁷(CH₂)_iR⁶, -SO₂(CH₂)_i(C₆-C₁₀ aryl), and -SO₂(CH₂)_i(5 to 10 membered heterocyclic), the -(CH₂)_q- and -(CH₂)_i- moieties of the said R⁵ groups optionally include a carbon-carbon double or triple bond, and the alkyl, aryl and heterocyclic moieties of the said

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30 R⁵ groups are unsubstituted or substituted with one or more substituents independently selected from halo, cyano, nitro, trifluoromethyl, azido, -OH, -C(O)R⁸, -C(O)OR⁸, -OC(O)R⁸,

-OC(O)OR⁸, -NR⁶C(O)R⁷, -C(O)NR⁶R⁷, -(CH₂)_iNR⁶R⁷, C₁-C₆ alkyl, C₃-C₁₀ cycloalkyl, -(CH₂)_i(C₆-C₁₀ aryl), -(CH₂)_i(5 to 10 membered heterocyclic), -(CH₂)_iO(CH₂)_qOR⁹, and -(CH₂)_iOR⁹;

each R⁶ and R⁷ is independently selected from H, OH, C₁-C₆ alkyl, C₃-C₁₀ cycloalkyl,
 5 -(CH₂)_i(C₆-C₁₀ aryl), -(CH₂)_i(5 to 10 membered heterocyclic), -(CH₂)_iO(CH₂)_qOR⁹,
 -(CH₂)_iCN(CH₂)_jOR⁹, -(CH₂)_iCN(CH₂)_jR⁹ and -(CH₂)_iOR⁹, and the alkyl, aryl and heterocyclic
 moieties of the said R⁶ and R⁷ groups are unsubstituted or substituted with one or more
 substituents independently selected from hydroxy, halo, cyano, nitro, trifluoromethyl, azido,
 -C(O)R⁸, -C(O)OR⁸, -CO(O)R⁸, -OC(O)OR⁸, -NR⁹C(O)R¹⁰, -C(O)NR⁹R¹⁰, -NR⁹R¹⁰, C₁-C₆ alkyl,
 10 -(CH₂)_i(C₆-C₁₀ aryl), -(CH₂)_i(5 to 10 membered heterocyclic), -(CH₂)_iO(CH₂)_qOR⁹, and
 -(CH₂)_iOR⁹, where when R⁶ and R⁷ are both attached to the same nitrogen, then R⁶ and R⁷
 are not both bonded to the nitrogen directly through an oxygen;

each R⁸ is independently selected from H, C₁-C₁₀ alkyl, C₃-C₁₀ cycloalkyl,
 -(CH₂)_i(C₆-C₁₀ aryl), and -(CH₂)_i(5 to 10 membered heterocyclic);
 15 t is an integer from 0 to 6; j is an integer from 0 to 2; q is an integer from 2 to 6;
 each R⁹ and R¹⁰ is independently selected from H, -OR⁶, C₁-C₆ alkyl, and
 C₃-C₁₀ cycloalkyl; and

each R¹² and R¹³ is independently selected from H, C₁-C₆ alkyl, C₃-C₁₀ cycloalkyl,
 -(CH₂)_i(C₃-C₁₀ cycloalkyl), -(CH₂)_i(C₆-C₁₀ aryl), -(CH₂)_i(5 to 10 membered heterocyclic),
 20 -(CH₂)_iO(CH₂)_qOR⁹, and -(CH₂)_iOR⁹, and the alkyl, aryl and heterocyclic moieties of the said
 R¹² and R¹³ groups are unsubstituted or substituted with one or more substituents
 independently selected from R⁵, or R¹² and R¹³ are taken together with the nitrogen to which
 they are attached to form a C₅-C₉ azabicyclic, aziridinyl, azetidiny, pyrrolidinyl, piperidyl,
 piperazinyl, morpholinyl, thiomorpholinyl, isoquinolinyl, or dihydroisoquinolinyl ring, wherein
 25 said C₅-C₉ azabicyclic, aziridinyl, azetidiny, pyrrolidinyl, piperidinyl, piperazinyl, morpholinyl,
 thiomorpholinyl, isoquinolinyl, or dihydroisoquinolinyl rings are unsubstituted or substituted
 with one or more R⁵ substituents, where R¹² and R¹³ are not both bonded to the nitrogen
 directly through an oxygen;

or prodrugs or metabolites thereof, or pharmaceutically acceptable salts or solvates of
 30 said compounds and said prodrugs and said metabolites.

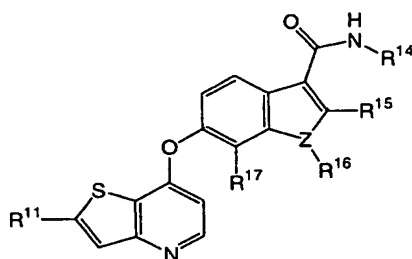
2. A compound, prodrug, metabolite, salt, or solvate according to claim 1, wherein R¹¹ is
 -(CH₂)_i(5 to 10 membered heterocyclic), -C(O)NR¹²R¹³, -(CH₂)_iNR¹²R¹³, -SO₂NR¹²R¹³ or
 -CO₂R¹².

3. A compound of claim 2, wherein R¹¹ is -(CH₂)_i(5 to 10 membered heterocyclic),
 35 -C(O)NR¹²R¹³, -SO₂NR¹²R¹³ or -CO₂R¹².

4. A compound of claim 3, wherein R¹¹ is -(CH₂)_i(5 to 10 membered heterocyclic) or
 -C(O)NR¹²R¹³.

5. A compound of claim 4, wherein R^{11} is $-C(O)NR^{12}R^{13}$, wherein R^{12} and R^{13} are independently selected from H, C_1 - C_6 alkyl, C_3 - C_{10} cycloalkyl, $-(CH_2)_i(C_3$ - C_{10} cycloalkyl), $-(CH_2)_i(C_6$ - C_{10} aryl), $-(CH_2)_i(5$ to 10 membered heterocyclic), $-(CH_2)_iO(CH_2)_qOR^9$, and $-(CH_2)_iOR^9$.
- 5 6. A compound of claim 5, wherein R^{11} is $-C(O)NR^{12}R^{13}$, and wherein R^{12} and R^{13} are taken together with the nitrogen to which they are attached to form a C_5 - C_9 azabicyclic, aziridinyl, azetidiny, pyrrolidinyl, piperidinyl, piperazinyl, morpholinyl, thiomorpholinyl, isoquinolinyl, or dihydroisoquinolinyl ring, wherein said C_5 - C_9 azabicyclic, aziridinyl, azetidiny, pyrrolidinyl, piperidinyl, piperazinyl, morpholinyl, thiomorpholinyl, isoquinolinyl, or dihydroisoquinolinyl ring is unsubstituted or substituted by 1 to 5 R^5 substituents.
- 10 7. A compound of claim 6, wherein R^{12} and R^{13} are taken together with the nitrogen to which they are attached to form a pyrrolidinyl, piperidinyl, piperazinyl, morpholinyl, thiomorpholinyl, isoquinolinyl, or dihydroisoquinolinyl ring, wherein said pyrrolidinyl, piperidinyl, piperazinyl, morpholinyl, thiomorpholinyl, isoquinolinyl, or dihydroisoquinolinyl ring is unsubstituted or substituted with 1 to 5 R^5 substituents.
- 15 8. A compound of claim 7, wherein R^{12} and R^{13} are taken together with the nitrogen to which they are attached to form a pyrrolidinyl, piperidinyl, piperazinyl, morpholinyl, or thiomorpholinyl ring, wherein said pyrrolidinyl, piperidinyl, piperazinyl, morpholinyl, or thiomorpholinyl ring is unsubstituted or substituted with 1 to 5 R^5 substituents.
- 20 9. A compound of claim 8, wherein R^{12} and R^{13} are taken together with the nitrogen to which they are attached to form a pyrrolidinyl or piperidinyl ring, wherein said pyrrolidinyl or piperidinyl ring is unsubstituted or substituted with 1 to 5 R^5 substituents.
- 25 10. A compound of claim 9, wherein R^{12} and R^{13} are taken together with the nitrogen to which they are attached to form a pyrrolidinyl ring, wherein said pyrrolidinyl is unsubstituted or substituted with 1 to 5 R^5 substituents.
11. A compound of claim 10, wherein R^{12} and R^{13} are taken together with the nitrogen to which they are attached to form a pyrrolidin-1-yl ring, wherein said pyrrolidin-1-yl ring is unsubstituted or substituted with 1 to 5 R^5 substituents.
- 30 12. A compound of claim 4, wherein R^{11} is a $-(CH_2)_i(5$ to 10 membered heterocyclic) group unsubstituted or substituted with 1 to 5 R^5 groups.
13. A compound of claim 12, wherein R^{11} is a $-(CH_2)_i(5$ -8 membered heterocyclic) group unsubstituted or substituted with 1 to 5 R^5 groups.
14. A compound of claim 13, wherein R^{11} is a $-(CH_2)_i(5$ or 6 membered heterocyclic) group is unsubstituted or substituted with 1 to 5 R^5 groups.
- 35 15. A compound of claim 14, wherein R^{11} is a $-(CH_2)_i(5$ membered heterocyclic) group unsubstituted or substituted with 1 to 5 R^5 groups.

16. A compound of claim 15, wherein R^{11} is a thiazolyl, unsubstituted or substituted by 1 to 5 R^5 groups.
17. A compound of claim 15, wherein R^{11} is an imidazolyl, unsubstituted or substituted by 1 to 5 R^5 groups.
- 5 18. A compound of claim 1, wherein R^{16} is a C_1 - C_6 alkyl group.
19. A compound of claim 18, wherein R^{16} is methyl.
20. A compound of claim 1, wherein R^{14} is methyl.
21. A compound represented by the formula II:



II

10 wherein:

Z is -O-, -S-, or -N-;

R^{14} is a C_1 - C_6 alkyl, C_1 - C_6 alkylamino, C_1 - C_6 alkylhydroxy, C_3 - C_{10} cycloalkylamino, or methylureido group;

R^{15} and R^{17} are independently H, halo, or a C_1 - C_6 alkyl group;

15 R^{16} is H or a C_1 - C_6 alkyl group when Z is -N- and R^{16} is absent when Z is -O- or -S-;

R^{11} is a heteroaryl group unsubstituted or substituted by one or more halo, cyano, nitro, trifluoromethoxy, trifluoromethyl, azido, $-C(O)R^8$, $-C(O)OR^8$, $-OC(O)R^8$, $-OC(O)OR^8$, $-NR^6C(O)R^7$, $-C(O)NR^6R^7$, $-NR^6R^7$, $-OR^9$, $-SO_2NR^6R^7$, C_1 - C_6 alkyl, C_3 - C_{10} cycloalkyl, $-(CH_2)_iO(CH_2)_qNR^6R^7$, $-(CH_2)_iO(CH_2)_qOR^9$, $-(CH_2)_iOR^9$, $-S(O)_i(C_1$ - C_6 alkyl), $-(CH_2)_i(C_6$ - C_{10} aryl), $-(CH_2)_i(5$ to 10 membered heterocyclic), $-C(O)(CH_2)_i(C_6$ - C_{10} aryl), $-(CH_2)_iO(CH_2)_j(C_6$ - C_{10} aryl), $-(CH_2)_iO(CH_2)_j(5$ to 10 membered heterocyclic), $-C(O)(CH_2)_i(5$ to 10 membered heterocyclic), $-(CH_2)_iNR^7(CH_2)_qNR^6R^7$, $-(CH_2)_iNR^7CH_2C(O)NR^6R^7$, $-(CH_2)_iNR^7(CH_2)_qNR^6C(O)R^8$, $-(CH_2)_iNR^7(CH_2)_iO(CH_2)_qOR^9$, $-(CH_2)_iNR^7(CH_2)_qS(O)_i(C_1$ - C_6 alkyl), $-(CH_2)_iNR^7$, $-(CH_2)_iR^6$, $-SO_2(CH_2)_i(C_6$ - C_{10} aryl), and $-SO_2(CH_2)_i(5$ to 10 membered heterocyclic), the $-(CH_2)_q$ - and $-(CH_2)_i$ - moieties of the said R^5 groups optionally include a carbon-carbon double or triple bond, and the alkyl, aryl and heterocyclic moieties of the said R^5 groups are unsubstituted or substituted with one or more substituents independently selected from halo, cyano, nitro, trifluoromethyl, azido, -OH, $-C(O)R^8$, $-C(O)OR^8$, $-OC(O)R^8$, $-OC(O)OR^8$, $-NR^6C(O)R^7$, $-C(O)NR^6R^7$, $-(CH_2)_iNR^6R^7$, C_1 - C_6 alkyl, C_3 - C_{10} cycloalkyl, $-(CH_2)_i(C_6$ - C_{10} aryl), $-(CH_2)_i(5$ to 10 membered heterocyclic), $-(CH_2)_iO(CH_2)_qOR^9$, and $-(CH_2)_iOR^9$;

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each R^6 and R^7 is independently selected from H, OH, C_1 - C_6 alkyl, C_3 - C_{10} cycloalkyl, $-(CH_2)_t(C_6-C_{10} \text{ aryl})$, $-(CH_2)_t(5 \text{ to } 10 \text{ membered heterocyclic})$, $-(CH_2)_tO(CH_2)_qOR^9$, $-(CH_2)_tCN(CH_2)_tOR^9$, $-(CH_2)_tCN(CH_2)_tR^9$ and $-(CH_2)_tOR^9$, and the alkyl, aryl and heterocyclic moieties of the said R^6 and R^7 groups are unsubstituted or substituted with one or more substituents independently selected from hydroxy, halo, cyano, nitro, trifluoromethyl, azido, $-C(O)R^8$, $-C(O)OR^8$, $-CO(O)R^8$, $-OC(O)OR^8$, $-NR^9C(O)R^{10}$, $-C(O)NR^9R^{10}$, $-NR^9R^{10}$, C_1 - C_6 alkyl, $-(CH_2)_t(C_6-C_{10} \text{ aryl})$, $-(CH_2)_t(5 \text{ to } 10 \text{ membered heterocyclic})$, $-(CH_2)_tO(CH_2)_qOR^9$, and $-(CH_2)_tOR^9$, where when R^6 and R^7 are both attached to the same nitrogen, then R^6 and R^7 are not both bonded to the nitrogen directly through an oxygen;

each R^8 is independently selected from H, C_1 - C_{10} alkyl, C_3 - C_{10} cycloalkyl, $-(CH_2)_t(C_6-C_{10} \text{ aryl})$, and $-(CH_2)_t(5 \text{ to } 10 \text{ membered heterocyclic})$;

each R^9 and R^{10} is independently selected from H, C_1 - C_6 alkyl, and C_3 - C_{10} cycloalkyl;

t is an integer from 0 to 6; j is an integer from 0 to 2; q is an integer from 2 to 6;

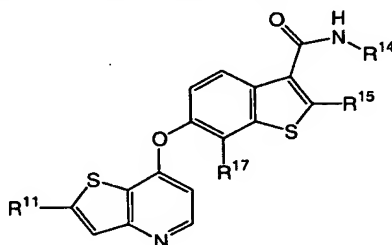
or prodrugs or metabolites thereof, pharmaceutically acceptable salts or solvates of said compounds, said prodrugs, or said metabolites.

22. A compound of claim 21, R^{16} is a C_1 - C_6 alkyl group.

23. A compound of claim 22, R^{16} is methyl.

24. A compound of claim 21, wherein R^{14} is methyl.

25. A compound represented by the formula III:



III

wherein:

R^{14} is a C_1 - C_6 alkyl, C_1 - C_6 alkylamino, C_1 - C_6 alkylhydroxy, C_3 - C_{10} cycloalkylamino, or methylureido group;

R^{15} and R^{17} are independently H, halo, or a C_1 - C_6 alkyl group; and

R^{11} is a heterocyclic or a heteroaryl group unsubstituted or substituted by one or more groups selected from $-C(O)OR^8$, C_1 - C_6 alkyl, and $-(CH_2)_tOR^9$;

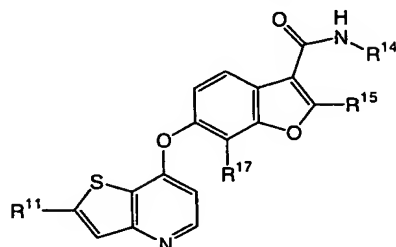
each R^8 is independently selected from H, C_1 - C_{10} alkyl, C_3 - C_{10} cycloalkyl, $-(CH_2)_t(C_6-C_{10} \text{ aryl})$, and $-(CH_2)_t(5 \text{ to } 10 \text{ membered heterocyclic})$;

each R^9 is independently selected from H, C_1 - C_6 alkyl, and C_3 - C_{10} cycloalkyl; and

t is an integer from 0 to 6; j is an integer from 0 to 2; q is an integer from 2 to 6;

or prodrugs or metabolites thereof, pharmaceutically acceptable salts or solvates of said compounds, said prodrugs, and said metabolites.

26. A compound of claim 25, wherein R^{14} is methyl.
 27. A compound represented by the formula IV:



IV

wherein:

- 5 R^{14} is a C_1 - C_6 alkyl, C_1 - C_6 alkylamino, C_1 - C_6 alkylhydroxy, C_3 - C_{10} cycloalkylamino, or methylureido group;

R^{15} and R^{17} are independently H, halo, or a C_1 - C_6 alkyl group;

R^{11} is a heterocyclic or a heteroaryl group unsubstituted or substituted by one or more groups selected from $-C(O)OR^8$, C_1 - C_6 alkyl, and $-(CH_2)_iOR^9$;

- 10 each R^8 is independently selected from H, C_1 - C_{10} alkyl, C_3 - C_{10} cycloalkyl, $-(CH_2)_j(C_6-C_{10}$ aryl), and $-(CH_2)_j(5 \text{ to } 10 \text{ membered heterocyclic})$;

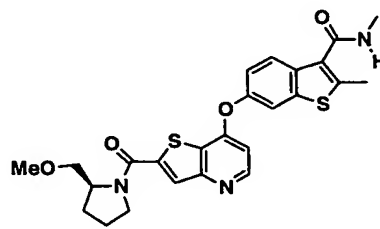
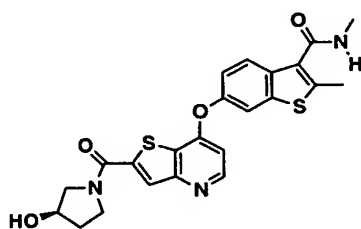
each R^9 is independently selected from H, C_1 - C_6 alkyl, and C_3 - C_{10} cycloalkyl; and

t is an integer from 0 to 6; j is an integer from 0 to 2; q is an integer from 2 to 6;

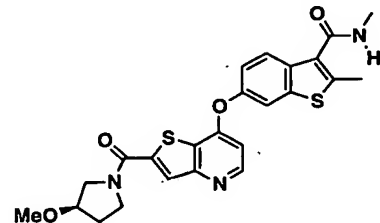
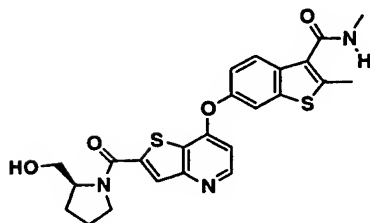
- or prodrugs or metabolites thereof, pharmaceutically acceptable salts or solvates of said compounds, said prodrugs, and said metabolites.

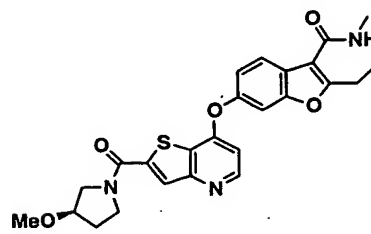
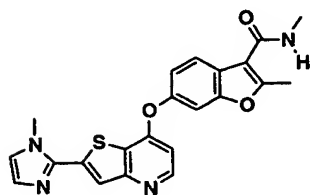
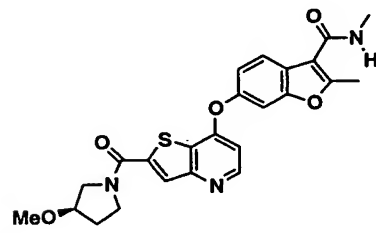
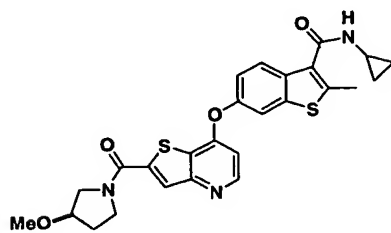
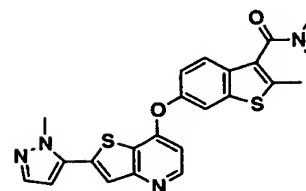
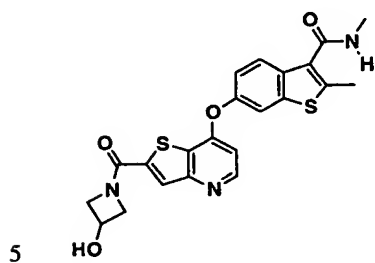
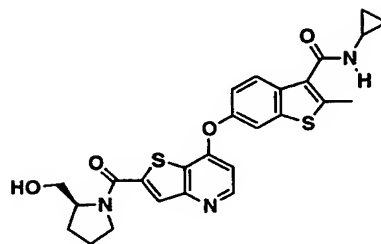
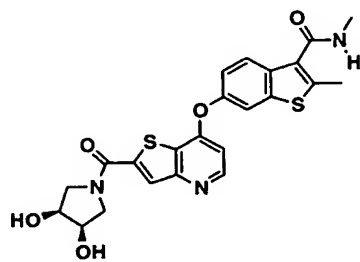
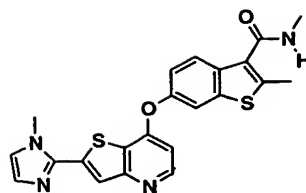
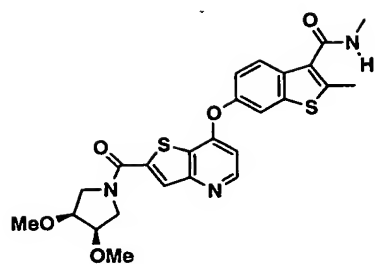
28. A compound of claim 27, wherein R^{14} is methyl.

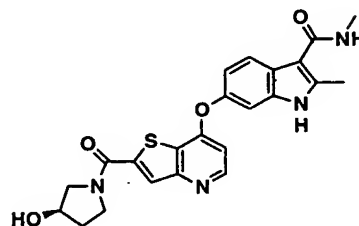
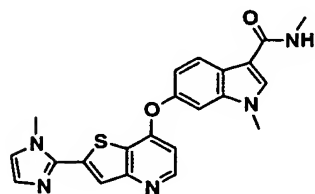
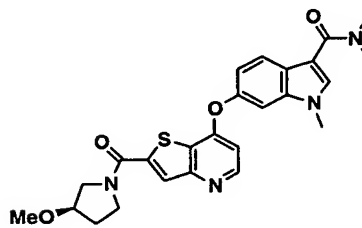
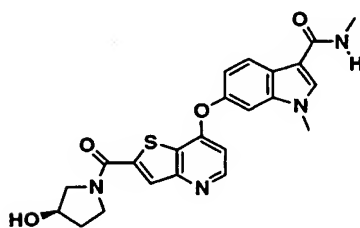
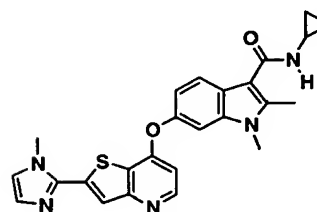
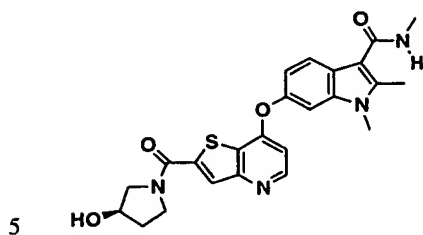
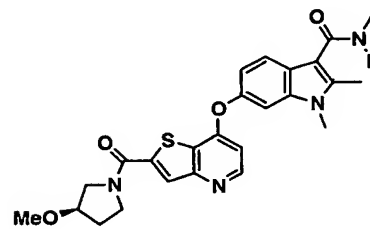
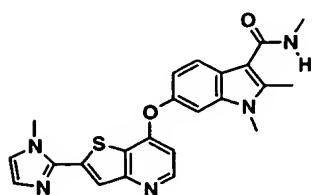
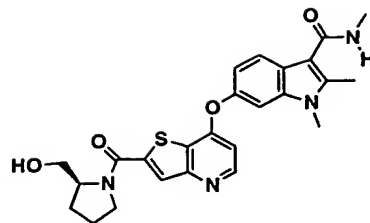
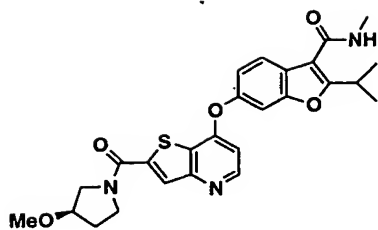
29. A compound of claim 1 wherein said compound is selected from the group consisting of:



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; and

or prodrugs or metabolites thereof, or pharmaceutically acceptable salts or solvates of said compounds, said prodrugs, and said metabolites.

30. A pharmaceutical composition for the treatment of a hyperproliferative disorder in a mammal comprising a therapeutically effective amount of a compound, prodrug, metabolite, salt or solvate of claim 1 and a pharmaceutically acceptable carrier.
- 5 31. The pharmaceutical composition of claim 30, wherein said hyperproliferative disorder is cancer.
32. The pharmaceutical composition of claim 31, wherein said cancer is brain, lung, kidney, renal, ovarian, ophthalmic, squamous cell, bladder, gastric, pancreatic, breast, head, neck, oesophageal, gynecological, prostate, colorectal or thyroid cancer.
- 10 33. The pharmaceutical composition of claim 30, wherein said hyperproliferative disorder is noncancerous.
34. The pharmaceutical composition of claim 33, wherein said hyperproliferative disorder is a benign hyperplasia of the skin or prostate.
- 15 35. A pharmaceutical composition for the treatment of a hyperproliferative disorder in a mammal comprising a therapeutically effective amount of a compound, prodrug, metabolite, salt or solvate of claim 1 in combination with an anti-tumor agent selected from the group consisting of mitotic inhibitors, alkylating agents, anti-metabolites, intercalating antibiotics, enzymes, topoisomerase inhibitors, biological response modifiers, anti-hormones, and anti-androgens, and a pharmaceutically acceptable carrier.
- 20 36. A pharmaceutical composition for the treatment of pancreatitis or kidney disease in a mammal comprising a therapeutically effective amount of a compound, prodrug, metabolite, salt or solvate of claim 1 and a pharmaceutically acceptable carrier.
37. A pharmaceutical composition for the prevention of blastocyte implantation in a mammal comprising a therapeutically effective amount of a compound, prodrug, metabolite, salt or solvate of claim 1 and a pharmaceutically acceptable carrier.
- 25 38. A pharmaceutical composition for treating a disease related to vasculogenesis or angiogenesis in a mammal comprising a therapeutically effective amount of a compound, prodrug, metabolite, salt or solvate of claim 1 and a pharmaceutically acceptable carrier.
- 30 39. The pharmaceutical composition of claim 38 wherein said disease is selected from the group consisting of tumor angiogenesis, chronic inflammatory disease, atherosclerosis, skin diseases, diabetes, diabetic retinopathy, retinopathy of prematurity, age-related macular degeneration, hemangioma, glioma, melanoma, Kaposi's sarcoma and ovarian, breast, lung, pancreatic, prostate, colon and epidermoid cancer.
- 35 40. A pharmaceutical composition for treating a disease related to vasculogenesis or angiogenesis in a mammal comprising a therapeutically effective amount of a compound, prodrug, metabolite, salt or solvate of claim 1, a therapeutically effective amount of a compound, prodrug, metabolite, salt or solvate of an antihypertensive agent, and a pharmaceutically acceptable carrier.

41. A method of treating a hyperproliferative disorder in a mammal comprising administering to said mammal a therapeutically effective amount of a compound, prodrug, metabolite, salt or solvate of claim 1.
42. The method of claim 41 wherein said hyperproliferative disorder is cancer.
- 5 43. The method of claim 42 wherein said cancer is brain, lung, ophthalmic, squamous cell, renal, kidney, ovarian, bladder, gastric, pancreatic, breast, head, neck, oesophageal, prostate, colorectal, gynecological or thyroid cancer.
44. The method of claim 41 wherein said hyperproliferative disorder is noncancerous.
45. The method of claim 44 wherein said hyperproliferative disorder is a benign hyperplasia
10 of the skin or prostate.
46. A method for the treatment of a hyperproliferative disorder in a mammal comprising administering to said mammal a therapeutically effective amount of a compound, prodrug, metabolite, salt or solvate of claim 1 in combination with an anti-tumor agent selected from the group consisting of mitotic inhibitors, alkylating agents, anti-metabolites, intercalating antibiotics,
15 growth factor inhibitors, cell cycle inhibitors, enzymes, topoisomerase inhibitors, biological response modifiers, anti-hormones, and anti-androgens.
47. A method of treating pancreatitis or kidney disease in a mammal comprising administering to said mammal a therapeutically effective amount of a compound, prodrug, metabolite, salt or solvate of claim 1.
- 20 48. A method of preventing blastocyte implantation in a mammal comprising administering to said mammal a therapeutically effective amount of a compound, prodrug, metabolite, salt or solvate of claim 1.
49. A method for treating a disease related to vasculogenesis or angiogenesis in a mammal comprising administering to said mammal a therapeutically effective amount of a
25 compound, prodrug, metabolite, salt or solvate of claim 1.
50. The method of claim 49 wherein said disease is selected from the group consisting of tumor angiogenesis, chronic inflammatory disease, atherosclerosis, skin diseases, diabetes, diabetic retinopathy, retinopathy of prematurity, age-related macular degeneration, hemangioma, glioma, melanoma, Kaposi's sarcoma and ovarian, breast, lung, pancreatic,
30 prostate, colon and epidermoid cancer.
51. A method for treating a disease related to vasculogenesis or angiogenesis in a mammal comprising administering to said mammal a therapeutically effective amount of a compound, prodrug, metabolite, salt or solvate of claim 1 in conjunction with a therapeutically effective amount of an anti-hypertensive agent.